

Supplementary Table S1. The main factors and the interactions of these factors affecting the secretion of LH by the porcine anterior pituitary cells during the estrous cycle (the results of multifactorial analysis of variance)

	FACTORS	F	p
Expt. No. 1	PHASE	$F_{3,512} = 2719.20$	0.00000
	STIMULATION OF CELLS	$F_{3,512} = 353.60$	0.00000
	VIS DOSE	$F_{3,512} = 2.70$	0.04245
	FK866	$F_{1,512} = 0.20$	0.62472
	PHASE*STIMULATION OF CELLS	$F_{9,512} = 105.00$	0.00000
	PHASE*VIS DOSE	$F_{9,512} = 12.90$	0.00000
	STIMULATION OF CELLS*VIS DOSE	$F_{9,512} = 23.90$	0.00000
	PHASE*FK866	$F_{3,512} = 138.90$	0.00000
	STIMULATION OF CELLS*FK866	$F_{3,512} = 120.70$	0.00000
	VIS DOSE*FK866	$F_{3,512} = 1.60$	0.18921
	PHASE*STIMULATION OF CELLS*VIS DOSE	$F_{27,512} = 9.70$	0.00000
	PHASE*STIMULATION OF CELLS*FK866	$F_{9,512} = 32.00$	0.00000
	PHASE*VIS DOSE*FK866	$F_{9,512} = 17.30$	0.00000
	STIMULATION OF CELLS*VIS DOSE*FK866	$F_{9,512} = 18.50$	0.00000
	PHASE*STIMULATION OF CELLS*VIS DOSE*FK866	$F_{27,512} = 6.50$	0.00000

Supplementary Table S2. The main factors and the interactions of these factors affecting the secretion of LH by the porcine anterior pituitary cells during the estrous cycle (the results of two-way analysis of variance)

		PHASE OF THE ESTROUS CYCLE	FACTORS	F	p	
Expt. No. 1	LH	basal secretion	early-luteal phase	VIS	$F_{3,32} = 8.67$	0.00024
				FK866	$F_{1,32} = 22.06$	0.00005
				VIS*FK866	$F_{3,32} = 5.33$	0.00431
			mid-luteal phase	VIS	$F_{3,32} = 6.64$	0.00129
				FK866	$F_{1,32} = 34.48$	0.00000
				VIS*FK866	$F_{3,32} = 8.09$	0.00038
			late-luteal phase	VIS	$F_{3,32} = 2.18$	0.10974
				FK866	$F_{1,32} = 14.53$	0.00059
				VIS*FK866	$F_{3,32} = 2.74$	0.05957
			follicular phase	VIS	$F_{3,32} = 7.03$	0.00092
				FK866	$F_{1,32} = 71.84$	0.00000
				VIS*FK866	$F_{3,32} = 8.99$	0.00018
GnRH-simulated secretion			early-luteal phase	VIS	$F_{3,32} = 39.26$	0.00000
				FK866	$F_{1,32} = 180.39$	0.00000
				VIS*FK866	$F_{3,32} = 19.68$	0.00000
			mid-luteal phase	VIS	$F_{3,32} = 13.12$	0.00001
				FK866	$F_{1,32} = 14.61$	0.00058
				VIS*FK866	$F_{3,32} = 6.63$	0.00131
			late-luteal phase	VIS	$F_{3,32} = 13.87$	0.00001
				FK866	$F_{1,32} = 95.55$	0.00000
				VIS*FK866	$F_{3,32} = 11.41$	0.00003
			follicular phase	VIS	$F_{3,32} = 6.57$	0.00138
				FK866	$F_{1,32} = 66.49$	0.00000
				VIS*FK866	$F_{3,32} = 9.20$	0.00016
INS-stimulated secretion			early-luteal phase	VIS	$F_{3,32} = 6.95$	0.00099
				FK866	$F_{1,32} = 27.21$	0.00001
				VIS*FK866	$F_{3,32} = 6.98$	0.00096
			mid-luteal phase	VIS	$F_{3,32} = 6.59$	0.00135
				FK866	$F_{1,32} = 24.86$	0.00002
				VIS*FK866	$F_{3,32} = 8.87$	0.00020
			late-luteal phase	VIS	$F_{3,32} = 3.78$	0.01980
				FK866	$F_{1,32} = 11.74$	0.00170
				VIS*FK866	$F_{3,32} = 4.56$	0.00907
			follicular phase	VIS	$F_{3,32} = 10.80$	0.00005
				FK866	$F_{1,32} = 111.97$	0.00000
				VIS*FK866	$F_{3,32} = 16.08$	0.00000
GnRH+INS-stimulated secretion			early-luteal phase	VIS	$F_{3,32} = 15.51$	0.00000
				FK866	$F_{1,32} = 97.86$	0.00000
				VIS*FK866	$F_{3,32} = 14.61$	0.00000
			mid-luteal phase	VIS	$F_{3,32} = 15.11$	0.00000
				FK866	$F_{1,32} = 98.02$	0.00000
				VIS*FK866	$F_{3,32} = 12.54$	0.00001
			late-luteal phase	VIS	$F_{3,32} = 5.84$	0.00268
				FK866	$F_{1,32} = 24.30$	0.00002
				VIS*FK866	$F_{3,32} = 3.63$	0.02315
			follicular phase	VIS	$F_{3,32} = 22.60$	0.00000
				FK866	$F_{1,32} = 137.49$	0.00000
				VIS*FK866	$F_{3,32} = 23.28$	0.00000

Supplementary Table S3. The main factors and the interactions of these factors affecting the secretion of FSH by the porcine anterior pituitary cells during the estrous cycle (the results of multifactorial analysis of variance)

	FACTORS	F	p
Expt. No. 1	FSH secretion	PHASE	$F_{3,512} = 1630.39$ 0.00000
		STIMULATION OF CELLS	$F_{3,512} = 139.91$ 0.00000
		VIS DOSE	$F_{3,512} = 34.66$ 0.00000
		FK866	$F_{1,512} = 0.29$ 0.83584
		PHASE*STIMULATION OF CELLS	$F_{9,512} = 7.93$ 0.00000
		PHASE*VIS DOSE	$F_{9,512} = 76.69$ 0.00000
		STIMULATION OF CELLS*VIS DOSE	$F_{9,512} = 9.66$ 0.00000
		PHASE*FK866	$F_{3,512} = 0.17$ 0.99705
		STIMULATION OF CELLS*FK866	$F_{3,512} = 0.13$ 0.94042
		VIS DOSE*FK866	$F_{3,512} = 1.10$ 0.34832
		PHASE*STIMULATION OF CELLS*VIS DOSE	$F_{27,512} = 10.73$ 0.00000
		PHASE*STIMULATION OF CELLS*FK866	$F_{9,512} = 0.98$ 0.45427
		PHASE*VIS DOSE*FK866	$F_{9,512} = 0.32$ 0.99969
		STIMULATION OF CELLS*VIS DOSE*FK866	$F_{9,512} = 0.06$ 0.99995
		PHASE*STIMULATION OF CELLS*VIS DOSE*FK866	$F_{27,512} = 0.21$ 1.00000

Supplementary Table S4. The main factors and the interactions of these factors affecting the secretion of FSH by the porcine anterior pituitary cells during the estrous cycle (the results of two-way analysis of variance)

	PHASE OF THE ESTROUS CYCLE	FACTORS	F	p
Expt. No. 1	FSH	VIS	$F_{3,32} = 5.04$	0.00569
		FK866	$F_{1,32} = 5.82$	0.02176
		VIS*FK866	$F_{3,32} = 3.36$	0.03076
		VIS	$F_{3,32} = 2.19$	0.10853
		FK866	$F_{1,32} = 0.02$	0.88447
		VIS*FK866	$F_{3,32} = 1.12$	0.35613
		VIS	$F_{3,32} = 0.37$	0.77751
		FK866	$F_{1,32} = 0.48$	0.49168
		VIS*FK866	$F_{3,32} = 0.25$	0.85770
		VIS	$F_{3,32} = 39.11$	0.00000
		FK866	$F_{1,32} = 2.16$	0.11170
		VIS*FK866	$F_{3,32} = 4.00$	0.01582
GnRH-simulated secretion	INS-stimulated secretion	VIS	$F_{3,32} = 1.72$	0.18299
		FK866	$F_{1,32} = 1.70$	0.20181
		VIS*FK866	$F_{3,32} = 1.04$	0.38658
		VIS	$F_{3,32} = 1.21$	0.32047
		FK866	$F_{1,32} = 2.31$	0.13801
		VIS*FK866	$F_{3,32} = 1.38$	0.26540
		VIS	$F_{3,32} = 0.21$	0.88617
		FK866	$F_{1,32} = 0.46$	0.50350
		VIS*FK866	$F_{3,32} = 0.04$	0.99097
		VIS	$F_{3,32} = 20.77$	0.00007
		FK866	$F_{1,32} = 2.49$	0.07774
		VIS*FK866	$F_{3,32} = 3.02$	0.04396
GnRH+INS-stimulated secretion	follicular phase	VIS	$F_{3,32} = 4.82$	0.03550
		FK866	$F_{1,32} = 0.44$	0.72388
		VIS*FK866	$F_{3,32} = 0.72$	0.54750
		VIS	$F_{3,32} = 1.60$	0.20766
		FK866	$F_{1,32} = 2.30$	0.13900
		VIS*FK866	$F_{3,32} = 1.11$	0.36042
		VIS	$F_{3,32} = 6.88$	0.01327
		FK866	$F_{1,32} = 0.07$	0.97788
		VIS*FK866	$F_{3,32} = 0.19$	0.90072
		VIS	$F_{3,32} = 16.41$	0.00030
		FK866	$F_{1,32} = 1.58$	0.21248
		VIS*FK866	$F_{3,32} = 1.29$	0.29609
	early-luteal phase	VIS	$F_{3,32} = 15.45$	0.00043
		FK866	$F_{1,32} = 1.86$	0.15617
		VIS*FK866	$F_{3,32} = 2.42$	0.08469
		VIS	$F_{3,32} = 2.41$	0.08388
		FK866	$F_{1,32} = 0.71$	0.40725
		VIS*FK866	$F_{3,32} = 1.37$	0.27029
		VIS	$F_{3,32} = 0.15$	0.92881
		FK866	$F_{1,32} = 3.73$	0.06243
		VIS*FK866	$F_{3,32} = 0.02$	0.99665
		VIS	$F_{3,32} = 4.05$	0.01509
		FK866	$F_{1,32} = 34.60$	0.00000
		VIS*FK866	$F_{3,32} = 4.21$	0.01289

Supplementary Table S5. The main factors and the interactions of these factors affecting the proliferation of the porcine anterior pituitary cells during the estrous cycle (the results of multifactorial analysis of variance)

	FACTORS	F	p
<i>Expt. No. 2</i>	PHASE	$F_{3,128} = 9.90$	0,00001
	VIS DOSE	$F_{3,128} = 68.60$	0,00000
	FK866	$F_{1,128} = 2.00$	0,11893
	PHASE*VIS DOSE	$F_{9,128} = 6.60$	0,00035
	PHASE*FK866	$F_{3,128} = 0.70$	0,57493
	VIS DOSE*FK866	$F_{3,128} = 0.10$	0,99953
	PHASE*VIS DOSE*FK866	$F_{9,128} = 0.10$	0,99974

Supplementary Table S6. The main factors and the interactions of these factors affecting the proliferation of the porcine anterior pituitary cells during the estrous cycle (the results of two-way analysis of variance)

PHASE OF THE ESTROUS CYCLE	FACTORS	F	p
early-luteal phase	VIS	$F_{3,32} = 22.27$	0.00005
	FK866	$F_{1,32} = 2.26$	0.10066
	VIS*FK866	$F_{3,32} = 2.25$	0.10021
mid-luteal phase	VIS	$F_{3,32} = 14.37$	0.00063
	FK866	$F_{1,32} = 2.23$	0.10321
	VIS*FK866	$F_{3,32} = 1.66$	0.19497
Expt. No. 2	VIS	$F_{3,32} = 10.54$	0.00274
	FK866	$F_{1,32} = 2.57$	0.07150
	VIS*FK866	$F_{3,32} = 1.19$	0.33033
follicular phase	VIS	$F_{3,32} = 23.54$	0.00003
	FK866	$F_{1,32} = 2.35$	0.30809
	VIS*FK866	$F_{3,32} = 1.68$	0.19134

Supplementary Table S7. The main factors and the interactions of these factors affecting the apoptosis process in the porcine anterior pituitary cells (the results of two-way analysis of variance)

APOPTOSIS	FACTORS	F	p
the early phase	VIS	$F_{3,32} = 0.28$	0.83850
	FK866	$F_{1,32} = 0.08$	0.78116
	VIS*FK866	$F_{3,32} = 1.63$	0.20123
the late phase	VIS	$F_{3,32} = 0.10$	0.96014
	FK866	$F_{1,32} = 0.14$	0.70632
	VIS*FK866	$F_{3,32} = 1.38$	0.26741
Expt. No. 3	VIS	$F_{3,32} = 0.18$	0.91039
	FK866	$F_{1,32} = 0.11$	0.73936
	VIS*FK866	$F_{3,32} = 1.56$	0.21742
dead cells	VIS	$F_{3,32} = 0.11$	0.95533
	FK866	$F_{1,32} = 0.80$	0.37754
	VIS*FK866	$F_{3,32} = 0.02$	0.99709

Supplementary Table S8. The main factors and the interactions of these factors affecting LH secretion by the porcine anterior pituitary cells after treatment with inhibitors of the INSR, AKT/PI3K, MAPK/ERK1/2, and AMPK signaling pathways (the results of multifactorial analysis of variance)

	FACTORS	F	P
<i>Expt. No. 5</i>	VIS	$F_{1,39} = 1.27$	0.26687
	S961	$F_{1,39} = 0.61$	0.43938
	S961*VIS	$F_{3,39} = 0.09$	0.76708
	LY294002	$F_{1,39} = 0.44$	0.51135
	LY294002*VIS	$F_{3,39} = 0.07$	0.79725
	U0126	$F_{1,39} = 0.83$	0.48781
	U0126*VIS	$F_{1,39} = 0.78$	0.51453
	DMPH	$F_{3,39} = 0.58$	0.63023
	DMPH*VIS	$F_{3,39} = 0.69$	0.56392

Supplementary Table S9. The main factors and the interactions of these factors affecting FSH secretion by the porcine anterior pituitary cells after treatment with inhibitors of the INSR, AKT/PI3K, MAPK/ERK1/2, and AMPK signaling pathways (the results of multifactorial analysis of variance)

	FACTORS	F	p
<i>Expt. No. 5</i>	VIS	$F_{1,39} = 4.93$	0.03218
	S961	$F_{1,39} = 3.74$	0.06011
	S961*VIS	$F_{3,39} = 5.07$	0.00455
	LY294002	$F_{1,39} = 0.33$	0.57180
	LY294002*VIS	$F_{3,39} = 2.80$	0.05236
	U0126	$F_{1,39} = 0.04$	0.83670
	U0126*VIS	$F_{3,39} = 4.30$	0.04457
	DMPH	$F_{1,39} = 2.46$	0.07677
	DMPH*VIS	$F_{3,39} = 4.93$	0.00522

The purity of the anterior lobe collection is assured by the fact that the anterior and posterior lobes of the porcine pituitary are morphologically distinct tissues, as confirmed by microscopic examination:

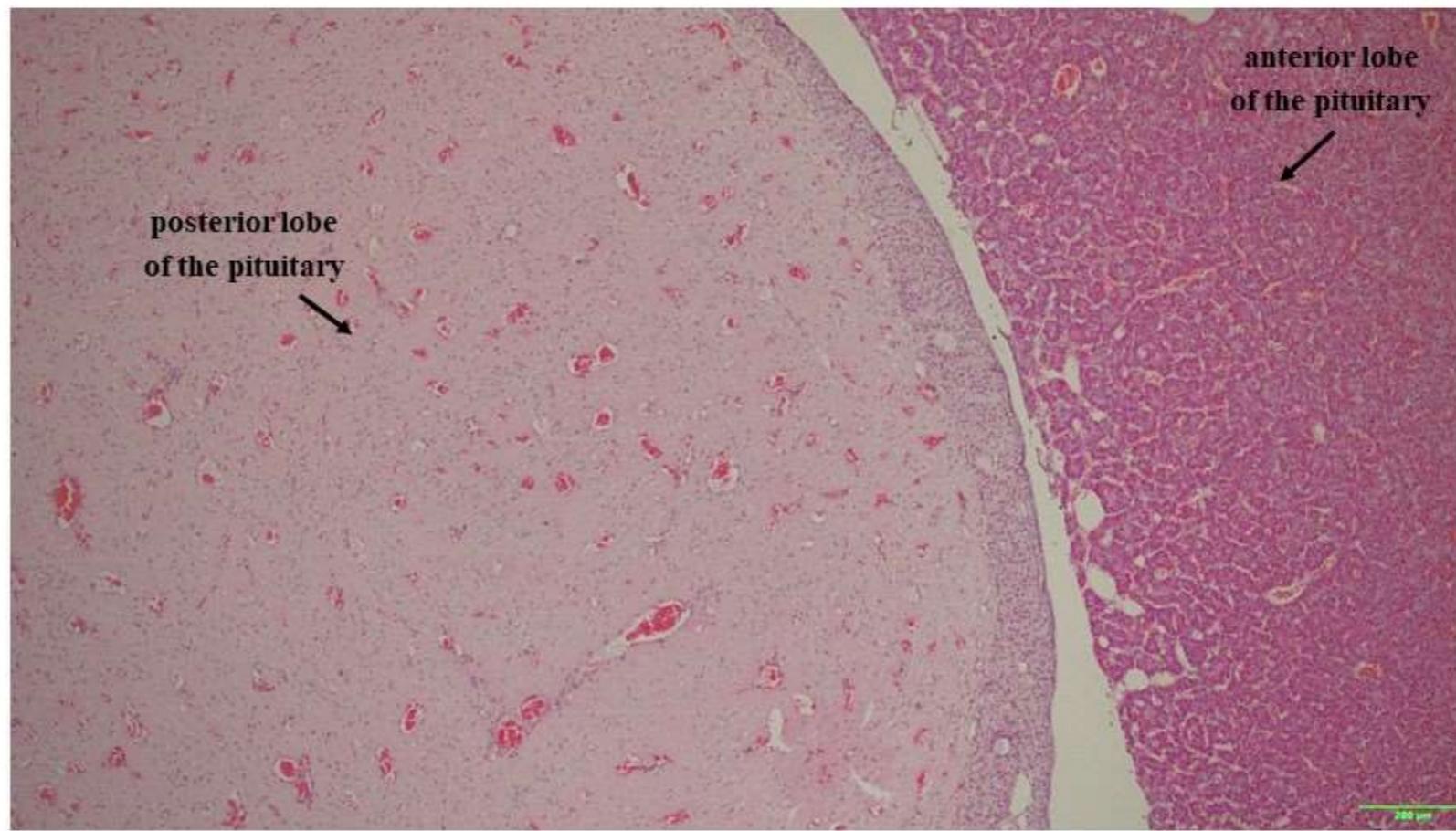
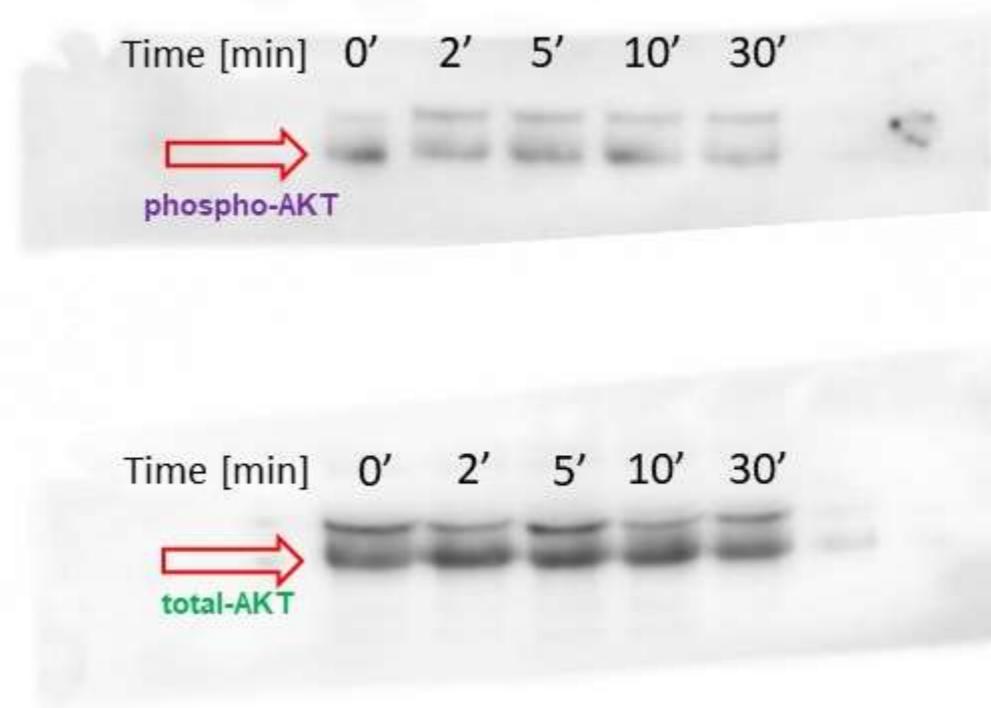
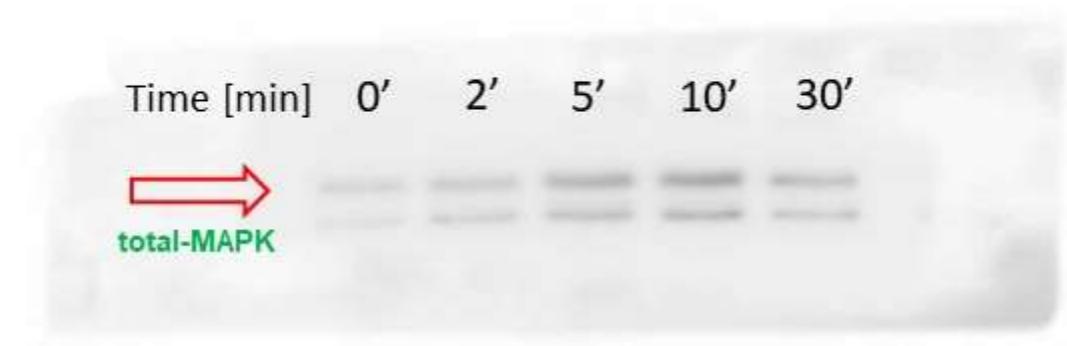
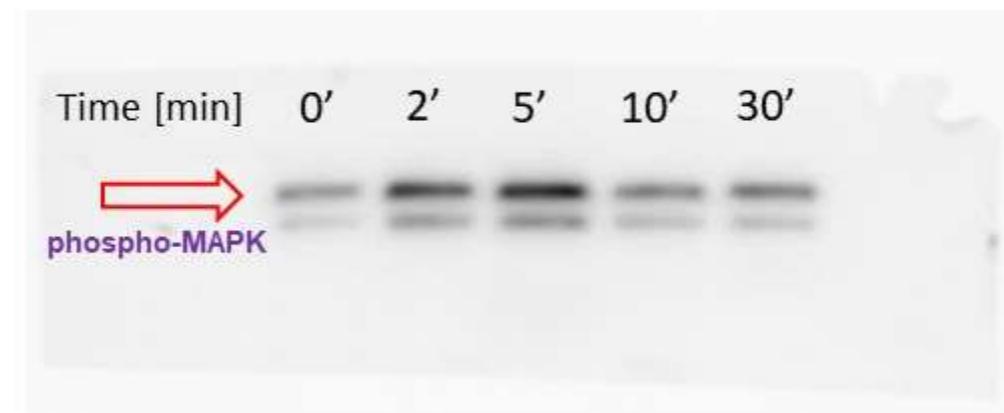


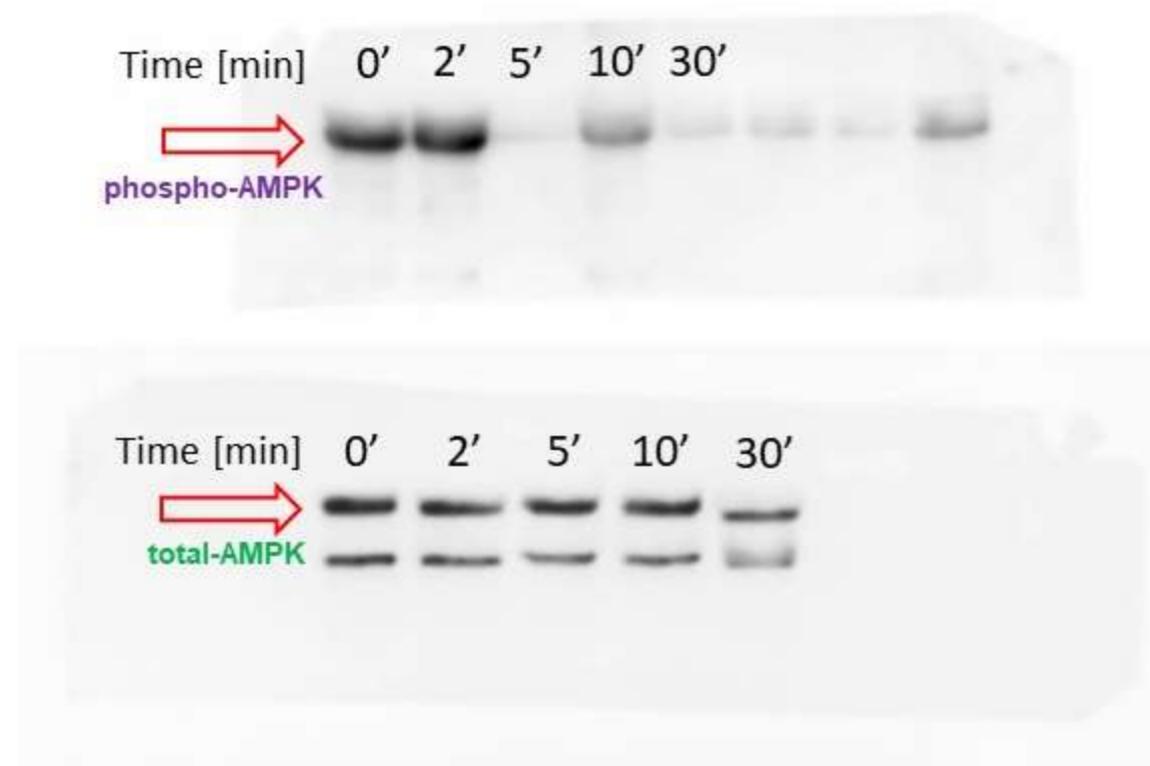
Figure shows the anterior and posterior lobes of the normal pituitary gland of the pig, x4.
Supplementary Figure S1. H&E staining of anterior and posterior pituitary.



Supplementary Figure S2. The in vitro effect of visfatin on the activation of the AKT/PI3K signalling pathways in the porcine anterior pituitary cells.



Supplementary Figure S3. The in vitro effect of visfatin on the activation of the MAPK/ERK1/2 signalling pathways in the porcine anterior pituitary cells.



Supplementary Figure S4. The in vitro effect of visfatin on the activation of the AMPK signalling pathways in the porcine anterior pituitary cells.